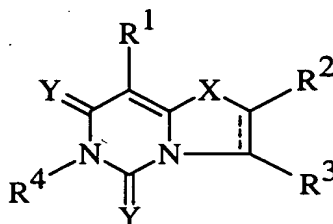


ABSTRACT

Selective MMP-13 inhibitors are bicyclic pyrimidines of the Formula I



or a pharmaceutically acceptable salt thereof,

wherein R^1 is hydrogen or alkyl; R^2 , R^3 , and R^4 include hydrogen, halo, alkyl, $C\equiv C(CH_2)_m$ aryl, CO_2 alkyl, $CO_2(CH_2)_m$ aryl, $COHN$ alkyl, and

$CONH(CH_2)_m$ aryl; and X is O, S, SO, SO_2 , CH_2 , $C=O$, $CHOH$, NH , or NR^5 .

A compound of Formula I, or a pharmaceutically acceptable salt thereof, is useful for treating cancer or arthritis.